

**DOCKET NO.:** ISIS-5243  
**Application No.:** Not Yet Assigned  
**Preliminary Amendment - First Action Not Yet Received**

**PATENT**

**Amendments to the Specification:**

Applicants respectfully request the application be amended, without prejudice, prior to examination as follows:

Please replace the paragraph beginning at page 1, line 3, with the following paragraph:

This application is a continuation of U.S. Patent Application Serial No. 10/269,291, filed October 11, 2002, which is a continuation of U.S. Serial No. 09/824,474, filed April 2, 2001, which is a continuation of SN 09/395,948 filed September 14, 1999, now U.S. Patent No. 6,211,350, which is a continuation of U.S. App. S.N. 08/692,909 filed July 31, 1996, now U.S. Patent No. 6,001,982, which is a division of U.S. S.N. 08/249,442 filed May 26, 1994, now U.S. Patent No. 5,571,902, which is a continuation-in-part of U.S. Serial No. 08/099,075 filed July 29, 1993. now U.S. Patent 5,614,621, the entirety of which is incorporated herein by reference.